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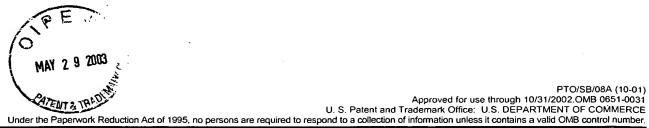
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Sut	ostitute for form 1449A/PTO			Complete if Known		
				Application Number	09/869,414	
11	VFORMATION	1 DI	SCLOSURE	Filing Date	June 27, 2001	
l s	STATEMENT I	3Y /	APPLICANT	First Named Inventor	Mark Gurney	
				Art Unit	1647	
	(use as many she	eets as	necessary)	Examiner Name	C. Nicholos NICHOLS	
Sheet	1	of	4	Attorney Docket Number	29915/6280M	

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	т —	Document Number		ATENT DOCUMENTS	Pages, Columns, Lines,
Examiner Initials*	Cite No.1	Number-Kind Code <sup>2</sup> (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Where Relevant Passages or Relevant Figures Appear
\$\frac{1}{2}\cdots\$	A1	5,424,205	6/13/95	Dovey et al.	
1	A2	5,593,846	1/14/97	Schenk et al.	
	A3	5,733,768	3/31/98	Dixon et al.	
	A4	5,744,346	4/28/98	Chrysler et al.	
	A5	5,750,349	5/12/98	Suzuki et al.	
	A6	5,766,846	6/16/98	Schlossmacher et al.	
	A7	5,837,672	11/17/98	Schenk et al.	
	A8	5,849,560	12/15/98	Abraham	
	A9	5,942,400	8/24/99	Anderson et al.	
	A10	6,025,180	2/15/00	Powell et al.	
	A11	5,455,169	10/3/95	Mullan	
	A12	5,795,963	8/18/98	Mullan	
		5,877,015	3/2/99	Hardy et al.	1
	A14	6,211,428	4/3/01	Singh et al.	
	A15	6,221,645	4/24/01	Chryster et al.	
	A16	6,245,884	6/12/01	Hook	
•		6,245,964	6/12/01	McLonlogue et al.	
	A18	60/141,363	<del> </del>	Lin et al.	
	<del>A19</del>	66/168,060		Lin et al.	
		60/178,368		Lin et al.	
		60/210,292		Hong et al.	
	A22	09/277,229	3	Citron et al.	/
		6,313,268		Hook	
	A24	60/177,836		Lin et al.	
	<del>A25</del>	60/119,571		Basi et al.	<del></del>
	A26	60/139,172		Anderson et al.	
	A27	60/114,408		Basi et al.	
	A28	09/404,578		Chrysler et al.	
	1	09/054,334		Anderson et al.	
١,	<b>V30</b>	09/730,329		Anderson et al.	
-		09/471,669		Anderson et al.	
	1	09/501,708		Anderson et al	
	L	09/723,722		Anderson et al.	
	1	09/724,566		Anderson et al.	
1		09/723,730		Anderson et al.	+
CW)	<del>A36</del>	09/724,571		Anderson et al.	<del></del>



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l s	STATEMENT I	BY A	APPLICANT	First Named Inventor	Mark Gurney	
				Art Unit	1647	
	(use as many sh	eets as	necessary)	Examiner Name	C. Nicholes NKHOLS	
Sheet	2	of	4	Attorney Docket Number	29915/6280M	

3	<u> </u>	A37	09/724,568		Andercen et al	
		<del>A38</del>	09/724,569		Anderson et al.	
		A39	6,319,689		Powell et al.	
		A40	6,162,630	12/19/00	Powell et al.	
		A41	6,319,689	11/20/01	Powell et al.	
		A42	6,358,725	3/19/02	Christie at al.	
		A43	6,361,975	3/26/02	Christie et al.	
			6,291,223	9/18/01	Christie et al.	
0	8	A45	6,545,127	4/08/03	Tang et al.	

		FOREI	GN PATENT	DOCUMENTS		
Examiner Initials*	Cite No.1	Foreign Patent Document  Country Code <sup>3</sup> -Number <sup>4</sup> -Kind Code <sup>5</sup> (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T⁵
202	B1	WO 96/31122	10/10/96	Dixon et al.		-,
	B2	WO 96/40885	12/19/96	Chrysler et al.		_
	B3	WO 98/13488	4/2/98	Dyrks et al.		_
	B4	WO 98/21589	5/22/98	Virginia Lee		_
	B5	EP 0848 062 A2	6/17/98	David J. Powell		_
	B6	WO 98/26059	6/18/98	Chrysler et al.		_
	B7	EP 0855 444 A2	7/29/99	David J. Powell		
$T^{-}$	B8	WO 99/34004	8/7/99	Klaus et al.		$\vdash$
	B9	WO 99/31236	6/24/99	Bougueleret et al.		=
	B10	WO 99/46281	9/16/99	Wood et al.		F
	B11	WO 99/64587	12/16/99	Rholam et al.		F
	B12	WO 00/23576	4/27/00	Hook		-
	B13	WO 00/47618	8/17/00	Anderson et al.		F
	B14	WO 00/58479	10/5/00	Citron et al.		_
	B15	WO 00/56871	9/28/00	Postina		_
	B16	WO 00/68266	11/16/00	Becker et al.		_
	B17	WO 00/69262	11/23/00	Zhong et al.		F
	B18	WO 01/00663	1/4/01	Tang et al.		
	B19	WO 01/00665	1/4/01	Tang et al.		L
	B20	WO 01/29563	4/26/01	Christie et al.		F
	B21	WO 01/31054	5/3/01	Christie et al.		F
<b>V</b>	B22	WO 01/36600	5/25/01	Zhu et al.		
c gr	<b>B</b> 23	WO 01/38487	5/31/01	Zhu et al.		F
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	-			Application Number	09/869,414	
١N	<b>VFORMATIC</b>	N DIS	CLOSURE	Filing Date	June 27, 2001	
S	STATEMENT	BY A	PPLICANT	First Named Inventor	Mark Gurney	
				Art Unit	1647	
	(use as many	sheets as n	ecessary)	Examiner Name	C. Nichelee NICHOLS	
Sheet	3	of	4	Attorney Docket Number	29915/6280M	

<sup>\*</sup>EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant

<sup>&</sup>lt;sup>1</sup> Applicant's unique citation designation number (optional). <sup>2</sup> See attached Kinds Codes of USPTO Patent Documents at <a href="www.uspto.gov">www.uspto.gov</a> or MPEP 901.04. <sup>3</sup> Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup> For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the application number of the patent document. <sup>5</sup> Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup> Applicant is to place a check mark here if English language Translation is attached.

		OTHER PRIOR ART – NON PATENT LITERATURE DOCUMENTS					
Examiner Initials	Cite No.1	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T <sup>2</sup>				
age	C1	CHYUNG et al. Novel β-Secretase Cleavage of β-Amyloid Precursor Protei in the Endoplasmic Reticulum/Intermediate Compartment of NT2N Cells, <i>Journal of Cell Biology</i> , 138: 671-680 (August 11, 1997).					
	C2	EVIN et al., Alzheimer's disease amyloid precursor protein (AβPP): proteolytic processing, secretases and βA4 amyloid production, <i>Amyloid; Int. J. Exp. Clin. Invest.</i> 1: 263-280 (September 8, 1994).					
	C3	HAASS et al., Amyloid β-peptide is Produced by Cultured Cells During Normal Metabolism, <i>Nature</i> , 359: 322-325 (September 24, 1992).					
	C4	HAASS et al., β-Amyloid Peptide and 3-kDa Fragment are Derived by Distinct Cellular Mechanisms, <i>Journal of Biochemistry</i> , 268: 3021-3024 (February 15, 1993).					
	C5	HAASS et al., The Swedish Mutation Causes Early-Onset Alzheimer's Disease by β- Secretase Cleavage Within the Secretory Pathway, <i>Nature Medicine</i> , 12: 1291-1296 (December 1995).					
	C6	HIROSAWA et al., Characterization of cDNA Clones Selected by the GeneMark Analysis from Size-Fractionated cDNA Libraries From Human Brain, DNA Res., 6(5): 329-336 (October 29, 1999).	_				
	C7	HUSSAIN et al., Identification of a Novel Aspartic Protease (Asp 2) as β-Secretase, <i>Molecular and Cellular Neuroscience</i> , 14: 419-427 (1999).	-				
	C8	KANG et al., The Precursor of Alzheimer's Disease Amyloid A4 Protein Resembles a Cell-Surface Receptor, <i>Nature</i> , 325: 733-736 (February 19, 1987).	-				
	C9	KITAGUCHI et al., Novel Precursor of Alzheimer's Disease Amyloid Protein Shows Protease Inhibitory Activity, <i>Nature</i> , 331: 530-532 (February 11, 1988).	_				
	C10	KNOPS et al., Cell-type and Amyloid Precursor Protein-type Specific Inhibition of Aβ Release by Bafilomycin A1, a Selective Inhibitor of Vacuolar ATPases, <i>Journal of Biological Chemistry</i> , 270: 2419-2422 (February 10, 1995).					
J	C11	KOO and SQUAZZO Evidence that Production and Release of Amyloid β-Protein Involves the Endocytic Pathway, <i>Journal of Biological Chemistry</i> , 269: 17386-17389 (July 1, 1994).					
(30)	C12	PONTE et al., A New A4 Amyloid mRNA Contains a Domain Homologous to Serine Proteinase Inhibitors, <i>Nature</i> , 331: 525-527 (February 11, 1988).					

C21

C22

(August 1992).

PTO/SB/08B (10-01)
Approved for use through 10/31/2002.OMB 0651-0031
U. S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

Complete if Known

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Sub	ostitute for fo	orm 1449B/PTO			Complete ii raiowii				
					Application Number	09/088,314 CA 1869414			
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S	TATE	MENT B	ΥA	PPLICANT	First Named Inventor	Mark Gurney			
					Group Art Unit	1647			
	(u:	se as many shee	ts as n	necessary)	Examiner Name	Sharon Turner NICHOC	$\overline{s}$		
Sheet		4	of	4	Attorney Docket Number	28341/6280NCP			
<b>X</b>	C13					Protein Cleaved at the 361: 260-263 (January 21,			
1	C14		SINHA et al., Purification and Cloning of Amyloid Precursor Protein β-Secretase from Human Brain, <i>Nature</i> , 402: 537-540 (December, 2 1999).						
	C15	SZECSI, T 210): 5-22			, Scand. J. Clin. L	.ab. Invest., 52 (suppl.			
	C16	Precursor	TANZI et al., Protease Inhibitor Domain Encoded by an Amyloid Protein Precursor mRNA Associated with Alzheimer's Disease, <i>Nature</i> , 331: 528-530 (February 11, 1988).						
	C17	VASSER et al., β-secretase Cleavage of Alzheimer's Amyloid Precursor Protein by the Transmembrane Aspartic Protease BACE, <i>Science</i> , 286 (5440): 735-41 (October 22, 1999).							
	C18	<del></del>							
	C19	ZHAO et al., β-Secretase Processing of the β-Amyloid Precursor Protein in Transgenic Mice Is Efficient in Neurons but Inefficient in Astrocytes, <i>Journal of Biological Chemistry</i> , 271: 31407-31411 (December 6, 1996).  PGT Search report for PCT/US 99/20881							
	<del>C20</del>	PGT Searce	<del>ch re</del>	port for PCT/US 9	9/20881				
	1004	B 41 11 1 A 5 1		A D (1)		411			

Examiner Signature	(1119)	Date 6/29/02
Signature	(N/MeZ)	Considered (5/21/05

area of Beta-Secretase, News 08/09/2000, www.elancorp.com.

MULLAN et al., A Pathogenic Mutation for Probable Alzheimer's Disease in

the APP Gene at the N-Terminus of β-Amyloid, Nature Genetics 1: 345-347,

Elan and Pharmacia form Alzheimer's disease research collaboration in the

\*EXAMINER: Initial inference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of his form with next communication to applicant.

<sup>&</sup>lt;sup>1</sup>Applicant's unique citation designation number (optional). <sup>2</sup>Applicant is to place a check mark here if English language Translation is attached.